PCT

WORLD INTELLECTUAL PROPERTY ORGANIZATION International Bureau



INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(51) International Patent Classification 4: A61K 39/12, 39/17, 39/215	A1	(11) International Publication Number: WO 86/ 00811
A61K 39/255, 39/39		(43	3) International Publication Date: 13 February 1986 (13.02.86)
(21) International Application Number: PCT/HU	J85/00()48	
(22) International Filing Date: 29 July 1985	(29.07.	85)	Published With international search report.
(31) Priority Application Number:	2919	/84	
(32) Priority Date: 30 July 1984	(30.07.	84)	
(33) Priority Country:	3	UE	•
(71)(72) Applicant and Inventor: CSATÁRY, Lás HU]; Deres u. 7, H-1124 Budapest (HU).	zló [H	נט/	
(74) Agent: PATENTBUREAU DANUBIA; E linszky ut 16, H-1368 Budapest (HU).	3ajcsy-2	Zsi-	
(81) Designated States: AT (European patent), AU ropean patent), BR, CH (European patent) ropean patent), DK, FI, FR (European patent), IT (European patent), IT (European patent), IT ropean patent), MC, NL (European patent) SE (European patent), SU, US.), DE (atent), P, LU (Eu- GB Eu-	
	:		

(54) Title: THERAPEUTIC PREPARATIONS FOR TREATMENT OF CERTAIN VIRAL AND OTHER DISEASES AND THE PREPARATION OF THE SAME

(57) Abstract

Therapeutic preparations against certain viral and other diseases of humans. The preparations are specified as formulas containing attenuated Newcastle disease virus, Marek's disease virus, avian bronchitis virus or avian encephalitis virus, mixed in carrier substance generally used in human pharmacology, if desired along with one or more optional potentiating preparations. Another object of the invention is the use of the above viruses in the treatment of virus diseases as well as malignant and benign tumours and collagen diseases.

FOR THE PURPOSES OF INFORMATION ONLY

Codes used to identify States party to the PCT on the front pages of pamphlets publishing international applications under the PCT.

ΑT	Austria	GA	Gabon	MR	Mauritania
ΑU	Australia	GB	United Kingdom	MW	Malawi
BB	Barbados	HU	Hungary	NL	Netherlands
BE	Belgium	II	Italy	NO	Norway
BG	Bulgaria	JP	Japan	RO	Romania
BR	Brazil	KP	Democratic People's Republic	SD	Sudan
CF	Central African Republic		of Korea	SE	Sweden
CG	Congo	KR	Republic of Korea	SN	Senegal
CH	Switzerland	LI	Liechtenstein	SU	Soviet Union
CM	Cameroon	LK	Sri Lanka	TD	Chad
DE	Germany, Federal Republic of	LU	Luxembourg	TG	Togo
DK	Denmark	MC	Monaco	US	United States of A

FR France ML Mali

~:===

WO 86/00811 PCT/HU85/00048

THEREPAUTIC PREPARATIONS FOR TREATMENT OF CERTAIN VIRAL AND OTHER DISEASES AND THE PREPARATION OF THE SAME

Technical field

5

10

The present invention relates to a set of therapeutic preparations (further on Preparation(s)) for the treatment of certain viral and other diseases, together with the complete technics of their production, including all materials and methods used.

The Preparations are effective against various pathological conditions caused by the Herpes group of viruses, and against certain malignant and benign tumours and collagen diseases as well.

Background art

Inactivated or live attenuated viruses have long been used all over the world for the specific prevention of several viral diseases of humans and animals. We have found that certain attenuated viral vaccines, prepared against some viral diseases of animals (Newcastle disease, Marek's disease, avian branchitis, avian encephalitis) had a distinct therapeutic effect against various human diseases of known viral or unknown etiology.

Disclosure of invention

Accordingly the object of the present invention is a

set of preparations containing attenuated (or killed)

Newcastle disease virus, Marek's disease virus, avian

bronchitis virus or avian encephalitis virus, mixed in a

carrier which is acceptable for use in humans. The present

invention relates also to all materials and methods used

in the production of all preparations.

ç

5

10

15

20

25

30

The Preparations have already been used with very good result in the conditions listed below:

- 1. The Preparation containing attenuated Newcastle disease virus was active against the following conditions:
- a) Infections caused by viruses of the Herpes group, such as Herpex simplex I and II; Herpes zooster; Cytomegalovirus; infectious mononucleoses (Epstein-Barr virus)
- b) Aphthostomatitis
 - c) Verruca vulgaris
 - d) Infantile pharyngeal papilloma
 - e) Human adenocarcinomas
 - f) Collagen diseases (e.g. nodous polyarteritis)
- g) Rheematic arthritis
 - 2. The Preparation containing attenuated Marek's disease virus acted against the following conditions:
 - a) Various human cancers (mainly adenocarcinomas)
 - b) Infections caused by viruses of the Herpes group as listed above (1.a)
 - 3. The Preparation containing attenuated avian bronchitis virus was active against the following conditions:
 - a) Various human cancers
 - b) Infections caused by the above listed (l.a) viruses of the Herpes goup.
 - 4. The Preparation containing attenuated avian encephalitis virus was active against the following conditions:
 - a) Various human cancers

While most vaccines used for specific disease prevention in humans contain killed (inactivated) virus, all Preparations contain live attenuated virus. The possible mechanism of action of the Preparations is thus either an interference with the

5

10

15

20

virulent viruses responsible for the actual disease, or a highly efficient aspecific stimulation of the natural protective mechanisms operative in the organism involved by disease.

The Preparations are produced by suspending live attenuated or inactivated, purified or highly purified and concentrated viruses in one of the carrier substances generally used in human pharmacology. The Preparations may be supplemented with potentiating drugs (e.g. tranquilizers).

The optimal form of delivery of the Preparations is in a freeze-dried form, in sealed ampoules, with the diluent supplied in a separate vial. In this form, the Preparations can be stored at room temperature for a practically unlimited time. Another forms of delivery are in solution, suppository, capsule, emulsion, suspension, ointment, etc. These should be stored in cold, for their activity may decay repidly at room temperature.

The dosage of the Preparations depends of their actual composition and form, on the patient's general condition, and on stage (severity) of the disease actually treated.

The Preparations are non-toxic, and have no side effect whatever.

The Preparations were tested in humans and animals, and showed a great therapeutic efficiency in both clinical and experimental trials.

Ä

ŗ

ì,

5

Claims

- 1. The rapeutic preparation for the treatment of viral diseases and against malignant and benign tumours and collagen diseases characterized by containing live attenuated viruses selected from the group consisting of Newcastle disease virus, Marek's disease virus, avian bronchitis virus and avian encephalitis virus.
- 2. The preparation according to claim 1 characterized by containing at least one carrier.
- 3. Use of Newcastle disease virus, Marek's disease virus, avian bronchitis virus and avian encephalitis virus in the treatment of viral diseases, against malignant and benign tumours and collagen diseases.
- 4. The use of viruses according to claim 3 in the treatment of diseases caused by the Herpes group of viruses.

INTERNATIONAL SEARCH REPORT

International Application No PCT/HU 85/00048

	EFFICATION OF SUBJECT MATTER (if several class		
	to international Patent Classification (IPC) or to both N		-
IPC	⁴ A 61 K 39/12, 39/17, 39	/215, 39/255, 39/39	£
II. FIELD	S SEARCHED .		•
	Minimum Docum	tentation Searched 7	
Classificati	on System	Classification Symbole	
Int.	C1. ⁴ A 61 K 39/12, 39/1	7, 39/215, 39/255	
		or than Minimum Documentation nts are included in the Fields Searched ⁸	
	·		
III. DOCL	MENTS CONSIDERED TO BE RELEVANT		
Category *	Citation of Document, 11 with Indication, where a	ppropriate, of the relevant passages 18	Relevant to Claim No. 13
х	Patent Abstracts of Japa cations, section C, vol. 30 March 1983, The Paten Government, page 76 C 18 58-116 422 (Midori Juji	7, no. 219, issued at Office Japanese 8, see Kokai-no.	(1,2)
х	US, A, 3 548 054 (R.E. B -15 December 1970 (15.12. line 10 - column 3, line	70), see column 2.	(1,2)
X	GB, A, 1 274 820 (TAKEDA KABUSHIKI KAISHA) 17 May claims 1,11-13.	YAKUHIN KOGYO 1972 (17.05.72), see	(1,2)
х	AT, B, 346 476 (M. SEVOI (10.11.78), see claims 1	AN) 10 November 1978	(1,2)
X	US, A, 3 674 861 (A.E. C 1972 (04.07.72), see abs examples 2,5,6; column 1	tract; claims 12.13:	(1,2)
	·	•	
"A" doc con "E" ear film "L" doc who cita "O" doc oth "P" doc late	at categories of cited documents; 18 sument defining the general state of the art which is no suidered to be of perticular relevance itier document but published on or after the internations ng date cument which may throw doubte on priority claim(s) of ich is cited to astablish the publication date of anothe altion or other special reason (as specified) cument referring to an oral disclosure, use, exhibition of lef means cument published prior to the international filing date but than the priority date claimed	"X" document of particular relever carnot be considered novel of involves in inventive step. "Y" document of particular releval cannot be considered novel of involves in inventive step. "Y" document of particular releval cannot be considered to involve in involves in involves in involves in involves in involves involves in involves	lict with the application but ide or theory underlying the idea; the claimed invention or cannot be considered to too; the claimed invention is an inventive step when the or more other such docu-obvious to a person same
	PIFICATION		
	e Actual Completion of the International Search ctober 1985 (11.10.85)	Date of Mailing of this international S 16 October 1985	
Internation	nal Searching Authority	Signature of Authorized Officer	·
AUST	RIAN PATENT OFFICE	Toloda	~ev

III. DOCU	MENTS CONSIDERED TO BE RELEVANT (CONTINUED FROM THE SECOND SHEET	ח
Category •	Citation of Document, with indication, where appropriate, of the relevant passages	Relevant to Claim No
A	EP, A1, O 086 025 (AKZO N.V.) 17 August 1983 (17.08.83), see claims 1,3.	(1,2)
A	US, A, 4 357 320 (P. APOUTOWEIL, M.M. KRAS- SELT) O2 November 1982 (O2.11.82), see abstract; claims 1-7.	(1,2)
A !	US, A, 4 049 794 (0C. STRAUB) 20 September 1977 (20.09.77), see abstract; column 1, line 65 - column 4, line 49.	(1,2)
A	DE, A, 2 120 929 (MERCK & CO INC.) 18 November 1971 (18.11.71), see claims 1,2.	(1,2)
A	DE, A, 1 492 083 (PHILIPS ROXANE INC.) 07 August 1969 (07.08.69), see page 1, line 1 - page 4, line 16.	(1,2)
A	AT, B, 343 797 (H. STICKL) 12 June 1978 (12.06.78), see claim 1.	(1,2)
A !	FR, A, 2 444 467 (GIST-BROCADES N.V.) 18 July 1980 (18.07.80), see claims 1,5,6; page 8, lines 5-24.	(1,2)
:	· :	
• :	· .	
,		
		t t 1
		!
•		•
		!
;		!
		<u>!</u> !
i		
}		
į		
1		•

· · · · · · · · · · · · · · · · · · ·	-
1	
·	
·	
ļ	
V. GBSERVATIONS WHERE CERTAIN CLAIMS WERE FOUND UNSEARCHABLE	
This international search report has not been established in respect of certain claims under Article 17(2) (a) for the fo	ollowing reasons:
1. Claim numbers3.1.4 because they relate to subject matter not required to be searched by this Authority, n	emely:
· ·	•
Methods for treatment of the human or animal body by	therapy -
see Article 17(2)a)i) and Rule 39.1, iv).	
see Altitle 1/(2/a/1) and Rule 33.1, 10/.	
2. Claim numbers, because they relate to parts of the international application that do not comply with the ments to such an extent that no meaningful international search can be carried out, searchcaby.	e prescribed require-
3. Claim numbers because they are dependent claims and are not drafted in accordance with the second a PCT Suis 6.4(a)	and third sentences of
PCT Rule 6.4(a).	and third sentances of
and the state of t	and third sentences of
PCT Rule 6.4(a).	and third sentances of
PCT Rule 6.4(a). VI. OBSERVATIONS WHERE UNITY OF INVENTION IS LACKING 2	and third sentences of
PCT Rule 6.4(a). VI. OBSERVATIONS WHERE UNITY OF INVENTION IS LACKING ²	and third sentances of
PCT Rule 6.4(a). VI. OBSERVATIONS WHERE UNITY OF INVENTION IS LACKING 2	and third sentences of
PCT Rule 6.4(a). VI. OSSERVATIONS WHERE UNITY OF INVENTION IS LACKING ² This International Searching Authority found multiple inventions in this international application as follows:	
PCT Rule 6.4(a). VI. OBSERVATIONS WHERE UNITY OF INVENTION IS LACKING 2	
PCT Rule 6.4(a). VI. OSSERVATIONS WHERE UNITY OF INVENTION IS LACKING ² This International Searching Authority found multiple inventions in this international application as follows: 1. As all required additional search fees were timely paid by the applicant, this international search report covers of the international application. 2. As only some of the required additional search fees were timely paid by the applicant, this international search	all searchaple claims
PCT Rule 6.4(a). VI. OBSERVATIONS WHERE UNITY OF INVENTION IS LACKING ² This International Searching Authority found multiple inventions in this international application as follows: 1. As all required additional search fees were timely paid by the applicant, this international search report covere of the International application.	all searchaple claims
PCT Rule 6.4(s). VI. OSSERVATIONS WHERE UNITY OF INVENTION IS LACKING ² This International Searching Authority found multiple inventions in this international application as follows: 1. As all required additional asserth fees were timely paid by the applicant, this international search report covers of the international application. 2. As only some of the required additional search fees were timely paid by the applicant, this international search	all searchable claims
PCT Rule 6.4(a). VI. OSSERVATIONS WHERE UNITY OF INVENTION IS LACKING ² This International Searching Authority found multiple inventions in this international application as follows: 1. As all required additional search fees were timely paid by the applicant, this international search report covers of the international application. 2. As only some of the required additional search fees were timely paid by the applicant, this international search those claims of the international application for which fees were paid, specifically claims: 1. No required additional search fees were timely paid by the applicant. Consequently, this international search	all searchaple claims
PCT Rule 6.4(a). VI. OSSERVATIONS WHERE UNITY OF INVENTION IS LACKING ² This International Searching Authority found multiple inventions in this international application as follows: 1. As all required additional search fees were timely paid by the applicant, this international search report covers of the International application. 2. As only some of the required additional search fees were timely paid by the applicant, this international search those claims of the international application for which fees were paid, specifically claims:	all searchaple claims
PCT Rule 6.4(a). VI. OSSERVATIONS WHERE UNITY OF INVENTION IS LACKING ² This International Searching Authority found multiple inventions in this international application as follows: 1. As all required additional search fees were timely paid by the applicant, this international search report covers of the international application. 2. As only some of the required additional search fees were timely paid by the applicant, this international search those claims of the international application for which fees were paid, specifically claims: 1. No required additional search fees were timely paid by the applicant. Consequently, this international search	all searchaple claims
PCT Rule 6.4(a). VI. OSSERVATIONS WHERE UNITY OF INVENTION IS LACKING 1 This International Searching Authority found multiple inventions in this international application as follows: 1. As all required additional search fees were timely paid by the applicant, this international search report covers of the international application. 2. As only some of the required additional search fees were timely paid by the applicant, this international search those claims of the international application for which fees were paid, specifically claims: 1. No required additional search fees were timely paid by the applicant. Consequently, this international search the invention first mentioned in the claims; it is covered by claim numbers:	all searchaple claims ch report covers only report is restricted to
PCT Rule 6.4(a). VI. OSSERVATIONS WHERE UNITY OF INVENTION IS LACKING ¹ This International Searching Authority found multiple inventions in this international application as follows: 1. As all required additional search fees were timely paid by the applicant, this international search report covers of the International application. 2. As only some of the required additional search fees were timely paid by the applicant, this international search those claims of the international application for which fees were paid, specifically claims: 1. No required additional search fees were timely paid by the applicant. Consequently, this international search the invention first mantioned in the claims; it is covered by claim numbers:	all searchaple claims ch report covers only report is restricted to
PCT Rule 6.4(a). VI. OBSERVATIONS WHERE UNITY OF INVENTION IS LACKING 2 This International Searching Authority found multiple inventions in this international application as follows: 1. As all required additional search fees were timely paid by the applicant, this international search report covers of the international application. 2. As only some of the required additional search fees were timely paid by the applicant, this international search those claims of the international application for which fees were paid, specifically claims: 1. No required additional search fees were timely paid by the applicant. Consequently, this international search the invention first mentioned in the claims; it is covered by claim numbers: 4. As all searchablis claims could be searched without effort justifying an additional fee, the international Searchablis claims could be searched without effort justifying an additional fee, the international Searchable claims could be searched without effort justifying an additional fee, the international Searchable claims could be searched without effort justifying an additional fee, the international Searchable claims could be searched without effort justifying an additional fee, the international Searchable claims could be searched without effort justifying an additional fee.	all searchaple claims ch report covers only report is restricted to

Form PCT/ISA/210 (aupplemental sheet (2)) (January 1985)

Anhang zum internationalen Recherchenbericht über die internationale Patentanmeldung

In diesem Anhang sind die Mitglieder der Patentfamilien der im obengenannten internationalen Recherchenbericht angeführten Patentdokumente angegeben. Diese Angaben dienen nur zur Unterrichtung und erfolgen ohne Gewähr.

Annex to the International Search Report on International Patent Application No. PCT/HU 85/00048

This Annex lists the patent family members relating to the patent documents cited in the above-mentioned International search report. The Austrian Patent Office is in no way liable for these particulars which are merely given for the purpose of information.

Annexe au rapport de recherche internationale relatif à la demande de brevet international n°.

La présente annexe indique les membres de la famille de brevets relatifs aux documents de brevets cités dans, le rapport de recherche internationale visé ci-dessus. Les renseignements fournis sont donnés à titre indicatif et n'engagent pas la responsabilité de l'Office autrichien des brevets.

Im Recherchenbericht angeführtes Patent- dokument Patent document cited in search report Document de brevet cité dans le rapport de recherche	Datum der Veröffentlichung Publication date Date de publication	Mitglied(er) der Patentfamilie Patent family member(s) Membre(s) de la famille de brevets	Datum der Veröffentlichung Publication date Date de publication
JP-A2-58 116 422 US-A - 3 548 054	30/03/1983 15/12/1970	None None	
GB-A - 1 274 820	17/05/1972	CA-A1- 962 593 DE-A -2 058 645 DE-C2-2 058 645 DK-B - 124 340 ES-A1- 385 968 FR-A1-2 073 436 FR-A5-2 073 436 FR-B1-2 073 436 NL-A -7 017 334 NL-B - 170 595 NL-C - 170 595	11/02/1975 09/06/1971 11/02/1982 09/10/1972 16/11/1973 01/10/1971 01/10/1971 30/08/1974 02/06/1971 01/07/1982 01/12/1982
AT-B - 346 476	10/11/1978	AR-A1- 201 878 AT-A - 6 002/74 AU-A1-70 962/74 BE-A1- 817 865 DE-A1-2 434 515 DK-A - 3 895/74 DK-B - 137 931 DK-C - 137 931 ES-A1- 428 370 FI-A - 219 974 FR-A1-2 301 268 FR-B1-2 301 268 GB-A -1 481 847 IE-B - 40 159 IL-A0- 45 181 IT-A -1 017 330 JP-A2-50 052 227 NL-A -7 409 842	24/04/1975 15/03/1978 08/01/1976 18/11/1974 27/03/1975 10/03/1975 05/06/1978 30/10/1978 16/09/1976 20/01/1975 17/09/1976 21/07/1978 03/08/1977 28/03/1979 22/10/1974 20/07/1977 09/05/1975
Fortsetzung siehe Bla	att 2	NO-A - 742 557 SE-A -7 409 411	21/01/1975 20/01/1975

				_
AT-B - 34	6 ⁻ 476	10/11/1978	US-A-3 981 771 ZA-A- 743 771	21/09/1976 30/07/1975
US-A - 3 67	4 861	04/07/1972	BE-A - 741 865 BG-D - 20 617 CA-A1- 966 779 DE-A - 1 957 747 DK-B - 131 545 DK-C - 131 609 ES-A1- 373 631 FR-A5- 2 023 537 FR-B1- 2 023 537 GB-A - 1 292 803 IL-A0- 33 318 IL-A1- 33 318 IT-A - 1 056 006 JP-B4-49 035 144 NL-A - 6 917 014 RO-P - 61 995	20/09/1974 20/05/1970
EP-A1-0 086	5 025	17/08/1983	AU-A1- 10 769/83 CA-A1- 1 189 450 DK-A0- 470/83 DK-A - 470/83 ES-A1- 519 551 ES-A5- 519 551 GB-A1- 2 114 438 IL-A0- 67 820 JP-A2-58 146 513 ZA-A - 830 430	25/06/1985 31/01/1983 06/08/1983 16/03/1984 13/04/1984 24/08/1983 01/09/1983
US-A -4 35	7 320	02/11/1982	US-E - 31 830 US-A - 4 500 638 US-A - 4 505 892 AR-A1- 222 263 AT-E - 4 510 CA-A2- 1 183 473 CA-A1- 1 184 115 CA-A2- 1 184 116 DK-A - 5 052/80 EP-A2- 0 030 063 EP-B1- 0 030 063 EP-B1- 0 030 063 EP-B1- 0 030 063 ES-A1- 497 253 JP-A2-56 092 823 NL-A - 7 908 683 PT-A - 72 07 PT-B - 72 07 ZA-A - 807 45 NL-A - 8 005 08 NL-A - 8 005 06	19/02/1985 19/03/1985 30/04/1981 15/09/1983 05/03/1985 19/03/1985 19/03/1985 19/03/1985 19/03/1985 19/03/1981 10/06/1981 10/06/1981 24/08/1983 01/11/1981 24/11/1981 27/07/1981 7 01/07/1981 7 01/07/1981 9 25/11/1981 3 01/04/1982

٠.

DE-A-2 120 929 18/11/1971 AT-B - 317 421 26/08/19 AT-B - 317 606 10/09/19 BE-A1- 766 492 16/09/19 BR-A0- 7 102 066 13/03/19 CA-A1- 934 295 25/09/19 CH-A - 565 863 29/08/19 CS-P - 191 868 31/07/19 DE-B2- 2 120 929 28/06/19	73 76 76 77 77 77 77 77 77 77 77 77 77 77
DK-B - 133 311 26/04/19 DK-C - 133 311 20/09/19 ES-A1- 390 698 01/07/19 FR-A5- 2 092 112 21/01/19 FR-B1- 2 092 112 06/09/19 GB-A - 1 316 168 09/05/19 GB-A - 1 317 710 23/05/19 HU-P - 162 835 28/04/19 IE-B - 35 138 26/11/19 Fortsetzung siehe Blatt 4 IL-A0- 36 329 26/05/1	974 971 973 975 975 976 976 976 972 973 973 973 975

DE-A-2 120 929	18/11/1971	IL-A1- 36 329 IT-A - 1 045 525 JP-B4-48 041 534 NL-A - 7 100 529 NL-B - 165 217 NL-C - 165 217 PL-P - 76 295 US-A - 3 642 574 YU-A - 625/71 YU-B - 35 785 ZA-A - 711 382 BR-A0- 7 401 519	25/04/1975 10/05/1980 07/12/1973 02/11/1971 15/10/1980 16/03/1981 28/02/1975 15/02/1972 31/10/1980 30/06/1981 29/11/1972 31/12/1974
DE-A-1 492 083	07/08/1969	DK-B - 116 232	22/12/1969
AT-B- 343 797	12/06/1978	AT-A - 5 989/75 BE-A1- 831 691 CA-A1- 1 057 195 CH-A - 614 627 DE-A1- 2 437 166 DE-B2- 2 437 166 DE-C3- 2 437 166 DK-A - 3 477/75 DK-B - 141 835 DK-C - 141 835 ES-A1- 439 966 FR-A1- 2 280 390 FR-B1- 2 280 390 FR-B1- 2 280 390 GB-A - 1 517 540 JP-A2-51 041 420 NL-A - 7 509 136 SE-A - 7 508 565 US-A - 4 053 582 ZA-A - 754 725	15/10/1977 17/11/1975 26/06/1979 14/12/1979 19/02/1976 27/01/1977 13/10/1977 02/02/1976 30/06/1980 10/11/1980 01/03/1977 27/02/1976 11/04/1980 12/07/1978 07/04/1976 03/02/1976 01/01/1977 30/06/1976
FR-A-2 444 467	18/07/1980	AT-E - 6 125 BE-A1- 880 777 CA-A1- 1 136 043 DK-A - 5 472/79 EP-A1- 0 013 449 EP-B1- 0 013 449 ES-A1- 487 062 ES-A5- 487 062 ES-A5- 494 598 ES-A5- 494 598 FR-A1- 2 444 467 FR-B1- 2 444 467 JP-A2-55 089 232 LU-A - 82 018 NL-A - 7 812 359 PT-A - 70 556 US-A - 4 515 777 ZA-A - 796 944 NL-A - 7 903 299	15/02/1984 20/06/1980 23/11/1982 21/06/1980 23/07/1980 08/02/1984 16/12/1980 16/01/1981 16/05/1981 16/06/1981 18/07/1980 02/11/1984 05/07/1980 23/07/1981 24/06/1980 01/12/1979 07/05/1985 26/11/1980 28/10/1980

This Page is Inserted by IFW Indexing and Scanning Operations and is not part of the Official Record

BEST AVAILABLE IMAGES

Defective images within this document are accurate representations of the original documents submitted by the applicant.

Defects in the images include but are not limited to the items checked:

☐ BLACK BORDERS
☐ IMAGE CUT OFF AT TOP, BOTTOM OR SIDES
☐ FADED TEXT OR DRAWING
BLURRED OR ILLEGIBLE TEXT OR DRAWING
☐ SKEWED/SLANTED IMAGES
☐ COLOR OR BLACK AND WHITE PHOTOGRAPHS
☐ GRAY SCALE DOCUMENTS
☐ LINES OR MARKS ON ORIGINAL DOCUMENT
☐ REFERENCE(S) OR EXHIBIT(S) SUBMITTED ARE POOR QUALITY
Потнев.

IMAGES ARE BEST AVAILABLE COPY.

As rescanning these documents will not correct the image problems checked, please do not report these problems to the IFW Image Problem Mailbox.